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<110> EVANS, RONALD M. <120> Novel steroid-activated nuclear receptors and uses therefor <130> SALK2270-5 <140> 10/081,555 <141> 2002-02-20 <150> 09/458,366 **RECEIVED** <151> 1999-12-09 <160> 09/227,718 AUG 3 0 2002 <161> 1999-01-08 <170> 09/005,286 TECH CENTER 1600/2900 <171> 1998-01-09 <180> 43 <190> PatentIn Ver. 2.1 **COPY OF PAPERS** ORIGINALLY FILED <210> 1 <211> 2068 <212> DNA <213> Homo sapiens <220> <221> CDS <222> (583)..(1884) <220> <221> modified base <222> (1263) <223> a, c, t, or g <400> 1 ggcacgagga gatctaggtt caaattaatg ttgcccctag tggtaaagga cagagaccct 60 cagactgatg aaatgcgctc agaattactt agacaaagcg gatatttgcc actctcttcc 120 ccttttcctg tgtttttgta gtgaagagac ctgaaagaaa aaagtaggga gaacataatg 180 agaacaaata cggtaatctc ttcatttgct agttcaagtg ctggacttgg gacttaggag 240 gggcaatgga gccgcttagt gcctacatct gacttggact gaaatatagg.tgagagacaa 300 gattgtctca tatccgggga aatcataacc tatgactagg acgggaagag gaagcactgc 360 ctttacttca gtgggaatct cggcctcagc ctgcaagcca agtgttcaca gtgagaaaag 420 caagagaata agctaatact cctgtcctga acaaggcagc ggctccttgg taaagctact 480

ccttgatcga tcctttgcac cggattgttc aaagtggacc ccaggggaga agtcggagca 540

aaqaacttac caccaagcag tccaagaggc ccagaagcaa ac ctg gag gtg aga

1

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Leu Glu Val Arg

ccc Pro 5	aaa Lys	gaa Glu	agc Ser	tgg Trp	aac Asn 10	cat His	gct Ala	gac Asp	ttt Phe	gta Val 15	cac His	tgt Cys	gag Glu	gac Asp	aca Thr 20	642
														gtc Val 35		690
														tat Tyr		738
		-	_		_	-		_	_	-				agg Arg		786
														gcc Ala		834
														ctg Leu		882
														gac Asp 115		930
														gaa Glu		978
														cag Gln		1026
atg Met	atg Met 150	atc Ile	agg Arg	gag Glu	ctg Leu	atg Met 155	gac Asp	gct Ala	cag Gln	atg Met	aaa Lys 160	acc Thr	ttt Phe	gac Asp	act Thr	1074
					_				_					agc Ser	_	1122
														gaa Glu 195		1170
gcc Ala	aag Lys	tgg Trp	agc Ser 200	cag Gln	gtc Val	cgg Arg	aaa Lys	gat Asp 205	ctg Leu	tgc Cys	tct Ser	ttg Leu	aag Lys 210	gtc Val	tct Ser	1218
														acn Xaa		1266
_	_	_		Arg					Ser	_	_			atg Met	_	1314

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														aaa Lys		1362
														ctg Leu 275		1410
														gtg Val		1458
							-	_			_			tgc Cys	_	1506
														atg Met		1554
														gag Glu		1602
														ggt Gly 355		1650
														att Ile		1698
ctg Leu	aag Lys	tcc Ser 375	tac Tyr	att Ile	gaa Glu	tgc Cys	aat Asn 380	cgg Arg	ccc Pro	cag Gln	cct Pro	gct Ala 385	cat His	agg Arg	ttc Phe	1746
_		_	_		_	_	_					_	_	atc Ile		1794
														ccc Pro		1842
_	_			_	cag Gln		_						-			1884
tga	gegge	ctg (cctt	gggt	ga ca	acctt	cgag	g agg	gcago	cag	acco	agag	gcc (ctcts	gagccg	1944
gcactcccgg gccaagacag atggacactg ccaagagccg acaatgccct gctggcctgt										2004						
ctccctaggg aattcctgct atgacagctg gctagcattc ctcaggaagg acatggggtg										2064						
ccc	2															2068

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<210> 2

<211> 434

<212> PRT

<213> Homo sapiens

<220>

<221> MOD RES

<222> (227)

<223> Threonine

<400> 2

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Leu Glu Val Arg Pro Lys Glu Ser Trp Asn His Ala Asp Phe Val His 1 5 10 15

Cys Glu Asp Thr Glu Ser Val Pro Gly Lys Pro Ser Val Asn Ala Asp 20 25 30

Glu Glu Val Gly Gly Pro Gln Ile Cys Arg Val Cys Gly Asp Lys Ala 35 40 45

Thr Gly Tyr His Phe Asn Val Met Thr Cys Glu Gly Cys Lys Gly Phe 50 60

Phe Arg Arg Ala Met Lys Arg Asn Ala Arg Leu Arg Cys Pro Phe Arg 65 70 75 80

Lys Gly Ala Cys Glu Ile Thr Arg Lys Thr Arg Arg Gln Cys Gln Ala 85 90 95

Cys Arg Leu Arg Lys Cys Leu Glu Ser Gly Met Lys Lys Glu Met Ile 100 105 110

Met Ser Asp Glu Ala Val Glu Glu Arg Arg Ala Leu Ile Lys Arg Lys 115 120 125

Lys Ser Glu Arg Thr Gly Thr Gln Pro Leu Gly Val Gln Gly Leu Thr 130 135 140

Glu Glu Gln Arg Met Met Ile Arg Glu Leu Met Asp Ala Gln Met Lys 145 150 155 160

Thr Phe Asp Thr Thr Phe Ser His Phe Lys Asn Phe Arg Leu Pro Gly
165 170 175

Val Leu Ser Ser Gly Cys Glu Leu Pro Glu Pro Leu Gln Ala Pro Ser 180 185 190

Arg Glu Glu Ala Ala Lys Trp Ser Gln Val Arg Lys Asp Leu Cys Ser 195 200 205

Leu Lys Val Ser Leu Gln Ala Ala Gly Gly Gly Trp Gln Cys Leu Glu 210 215 220

Leu Gln Xaa Pro Ser Arg Gln Trp Arg Lys Glu Ile Phe Ser Leu Leu 225 230 235 240

Pro His Met Ala Asp Met Ser Thr Tyr Met Phe Lys Gly Ile Ile Ser 245 250 255

Phe Ala Lys Val Ile Ser Tyr Phe Arg Asp Leu Pro Ile Glu Asp Gln 260 265 270

Ile Ser Leu Leu Lys Gly Ala Ala Phe Glu Leu Cys Gln Leu Arg Phe 275 280 285

Asn Thr Val Phe Asn Ala Glu Thr Gly Thr Trp Glu Cys Gly Arg Leu 290 295 300

Ser Tyr Cys Leu Glu Asp Thr Ala Gly Gly Phe Gln Gln Leu Leu Leu 305 310 315

Glu Pro Met Leu Lys Phe His Tyr Met Leu Lys Lys Leu Gln Leu His 325 330 335

Glu Glu Glu Tyr Val Leu Met Gln Ala Ile Ser Leu Phe Ser Pro Asp 340 345 350

Arg Pro Gly Val Leu Gln His Arg Val Val Asp Gln Leu Gln Glu Gln 355 360 365

Phe Ala Ile Thr Leu Lys Ser Tyr Ile Glu Cys Asn Arg Pro Gln Pro 370 375 380

Ala His Arg Phe Leu Phe Leu Lys Ile Met Ala Met Leu Thr Glu Leu 385 390 395 400

Arg Ser Ile Asn Ala Gln His Thr Gln Arg Leu Leu Arg Ile Gln Asp 405 410 415

Ile His Pro Phe Ala Thr Pro Leu Met Gln Glu Leu Phe Gly Ile Thr 420 425 430

Gly Ser

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<210> 3

<211> 25

<212> DNA

<213> Artificial Sequence

<220>

<223> Description of Artificial Sequence: Putative SXR response element from the steroid hydoxylase, rCYP3A1

<400> 3 tagacagttc atgaagttca tctac

25

<210> 4

<211> 25

<212> DNA

<213> Artificial Sequence

<220>

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<400> 4 taagcagttc ataaagttca tctac	25
<210> 5 <211> 25 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Putative SXR response element from the steroid hydoxylase, rUGT1A6	
<400> 5 actgtagttc ataaagttca catgg	25
<210> 6 <211> 26 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Putative SXR response element from the steroid hydoxylase, rbCYP2C1	
<400> 6 caatcagttc accaat	26
<210> 7 <211> 33 <212> DNA <213> Artificial Sequence	
<pre><220> <223> Description of Artificial Sequence: Putative SXR response element from the steroid hydoxylase, rP450R</pre>	
<400> 7 cacaggtgag ctgaggccag cagcaggtcg aaa	33
<210> 8 <211> 27 <212> DNA <213> Artificial Sequence	
<220> <223> Description of Artificial Sequence: Putative SXR response element from the steroid hydoxylase, rCYP2A1	
<400> 8 gtgcaggttc aactggaggt caacatg	27

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<210> 9
<211> 27
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<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Putative SXR
      response element from the steroid hydoxylase,
      rCYP2A2
<400> 9
                                                                    27
gtgctggttc aactggaggt cagtatg
<210> 10
<211> 27
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Putative SXR
      response element from the steroid hydoxylase,
      rCYP2C6
<400> 10
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<210> 11
<211> 27
<212> DNA
<213> Artificial Sequence
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<223> Description of Artificial Sequence: Putative SXR
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<212> DNA
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<223> Description of Artificial Sequence: Direct repeat
      with spacer of 0 nucleotides
<400> 12
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catagtcagg tcaaggtcag atcaac
<210> 13
<211> 27
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<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Direct repeat
      with spacer of 1 nucleotides
<400> 13
                                                                    27
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<210> 14
<211> 28
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Direct repeat
      with spacer of 2 nucleotides
<400> 14
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catagtcagg tcaataggtc agatcaac
<210> 15
<211> 29
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Direct repeat
     with spacer of 3 nucleotides
<400> 15
                                                                    29
catagtcagg tcatataggt cagatcaac
<210> 16
<211> 30
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Direct repeat
     with spacer of 4 nucleotides
<400> 16
catagtcagg tcatataagg tcagatcaac
                                                                    30
<210> 17
<211> 31
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Direct repeat
      with spacer of 5 nucleotides
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<400> 17 catagtcagg tcatatatag gtcagatcaa c	31
<210> 18	
<213> Artificial Sequence	
<pre><220> <223> Description of Artificial Sequence: Direct repeat with spacer of 6 nucleotides</pre>	
<400> 18 catagtcagg tcatatata ggtcaagatc aac	33
<210> 19 <211> 33 <212> DNA	
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210. 20	
<210> 20 <211> 36	
<212> DNA <213> Artificial Sequence	
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<400> 20 catagtcagg tcatatata ataaggtcag atcaac	36
<210> 21 <211> 41 <212> DNA	
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<220> <223> Description of Artificial Sequence: Direct repeat with spacer of 15 nucleotides	
<400> 21 catagtcagg tcatagtagt agtagtagag gtcagatcaa c	41
<210> 22 <211> 17 <212> DNA	

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<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Example of a
      response element suitable for practice of the
      invention method
<220>
<221> modified_base
<222> (7)..(11)
<223> This region may encompass 5, 4 or 3 nucleotides,
      independently selected from a, c, t or g
<400> 22
                                                                   17
agttcannnn ntgaact
<210> 23
<211> 18
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Example of a
      response element suitable for practice of the
      invention method
<220>
<221> modified base
<222> (7)..(12)
<223> a, c, t or g
<400> 23
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tgaactnnnn nnaggtca
<210> 24
<211> 18
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Synthetic
      oligonucleotide
<400> 24
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tgaactcaaa ggaggtca
<210> 25
<211> 18
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Inverted
      repeat response element with spacer of 0
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nucleotides

_	
<400> 25	
agcttaggtc atgaccta	18
0.00	
<210> 26	
<211> 19	
<212> DNA	•
<213> Artificial Sequence	
<220>	
<223> Description of Artificial Sequence: Inv	verted
repeat response element with spacer of	1
nucleotides	
<400> 26	
agcttaggtc agtgaccta	19
<210> 27	
<211> 20	· .
<212> DNA	
<213> Artificial Sequence	
.220	
<pre><220> <223> Description of Artificial Sequence: Inv</pre>	rartad
repeat response element with spacer of	
nucleotides	
nucleocides	
<400> 27	
agettaggte aegtgaeeta	20
	•
<210> 28	
<211> 21	
<212> DNA	•
<213> Artificial Sequence	
<220>	
<223> Description of Artificial Sequence: Inv	
repeat response element with spacer of nucleotides	3
nucleocides	
<400> 28	
agettaggte acagtgacet a	21
agoodaggoo adagogadoo a	
1	
<210> 29	
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<220>	_
<223> Description of Artificial Sequence: Inv	
repeat response element with spacer of	4
nucleotides	
4400- 20	
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Action (ANSTROLL CONTS).

ASSIST-

<210><211><211><212><213>	23			
<220> <223>	Description of Artificial Sequence: Inverted repeat response element with spacer of 5 nucleotides			
<400> agctta	30 aggtc acactgtgac cta			23
<210><211><211><212><213>	23			
<220> <223>	Description of Artificial Sequence: Inverted repeat response element with spacer of 6 nucleotides			
<400> agctti	31 gaac tcaaaggagg tca	٠		23
<210><211><211><212><213>	18		•	
<220> <223>	Description of Artificial Sequence: IR-M			
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<210><211><211><212><213>	33			
<400> tagaa	33 atga actcaaagga ggtcagtgag tgg			33
<210><211><212><213>	33			
<400> tagaat	34 catga actcaaagga ggtaagcaaa ggg			33

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<210> 35
<211> 32
<212> DNA
<213> Homo sapiens
<400> 35
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tagaatatta actcaatgga ggcagtgagt gg
<210> 36
<211> 25
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Synthetic
      oligonucleotide for PCR
<400> 36 '
                                                                    25
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<210> 37
<211> 25
<212> DNA
<213> Artificial Sequence
<223> Description of Artificial Sequence: Synthetic
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<211> 25
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                                                                   . 25
<210> 39
<211> 15
<212> DNA
<213> Artificial Sequence
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                                                                    15
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 <223> Description of Artificial Sequence: Synthetic
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 <223> Description of Artificial Sequence: Illustrative
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 <400> 42
                                                                     15
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 <210> 43
 <211> 15
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<213> Artificial Sequence
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 <223> Description of Artificial Sequence: Synthetic
       oligonucleotide
 <400> 43
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acaacttcat gaact